

SEQUENCE LISTING

<110> Colgan, Sean

<120> Compositions and Methods for Treating Hematologic Malignancies and Multiple Drug Resistance

<130> B0801/7233 (ERP)

<150> US 60/243,542

<151> 2000-10-26

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<170> PatentIn version 3.1

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His Cys Thr Gly His Ile His Val Tyr Asp Thr Asn Ser Asn Gln Pro
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Gln Cys Gly Tyr Lys Lys Pro Pro Met Thr Cys Leu Val Leu Ile Cys
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Glu Pro Ile Pro His Pro Ser Asn Ile Glu Ile Pro Leu Asp Ser Lys
225 230 235 240

Thr Phe Leu Ser Arg His Ser Leu Asp Met Lys Phe Ser Tyr Cys Asp
245 250 255

Glu Arg Ile Thr Glu Leu Met Gly Tyr Glu Pro Glu Glu Leu Leu Gly
260 265 270

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Arg Ser Ile Tyr Glu Tyr Tyr His Ala Leu Asp Ser Asp His Leu Thr
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Lys Thr His His Asp Met Phe Thr Lys Gly Gln Val Thr Thr Gly Gln
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Tyr Arg Met Leu Ala Lys Arg Gly Gly Tyr Val Trp Val Glu Thr Gln
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Ala Thr Val Ile Tyr Asn Thr Lys Asn Ser Gln Pro Gln Cys Ile Val
325 330 335

Cys Val Asn Tyr Val Val Ser Gly Ile Ile Gln His Asp Leu Ile Phe
340 345 350

Ser Leu Gln Gln Thr Glu Cys Val Leu Lys Pro Val Glu Ser Ser Asp
355 360 365

Met Lys Met Thr Gln Leu Phe Thr Lys Val Glu Ser Glu Asp Thr Ser
370 375 380

Ser Leu Phe Asp Lys Leu Lys Lys Glu Pro Asp Ala Leu Thr Leu Leu
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Ala Pro Ala Ala Gly Asp Thr Ile Ile Ser Leu Asp Phe Gly Ser Asn
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Asp Thr Glu Thr Asp Asp Gln Gln Leu Glu Glu Val Pro Leu Tyr Asn
420 425 430

Asp Val Met Leu Pro Ser Pro Asn Glu Lys Leu Gln Asn Ile Asn Leu
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Ala Met Ser Pro Leu Pro Thr Ala Glu Thr Pro Lys Pro Leu Arg Ser
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Ser Ala Asp Pro Ala Leu Asn Gln Glu Val Ala Leu Lys Leu Glu Pro
465 470 475 480

Asn Pro Glu Ser Leu Glu Leu Ser Phe Thr Met Pro Gln Ile Gln Asp
485 490 495

Gln Thr Pro Ser Pro Ser Asp Gly Ser Thr Arg Gln Ser Ser Pro Glu
500 505 510

Pro Asn Ser Pro Ser Glu Tyr Cys Phe Tyr Val Asp Ser Asp Met Val
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Asn Glu Phe Lys Leu Glu Leu Val Glu Lys Leu Phe Ala Glu Asp Thr
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Glu Ala Lys Asn Pro Phe Ser Thr Gln Asp Thr Asp Leu Asp Leu Glu
545 550 555 560

Met Leu Ala Pro Tyr Ile Pro Met Asp Asp Asp Phe Gln Leu Arg Ser
565 570 575

Phe Asp Gln Leu Ser Pro Leu Glu Ser Ser Ser Ala Ser Pro Glu Ser
580 585 590

Ala Ser Pro Gln Ser Thr Val Thr Val Phe Gln Gln Thr Gln Ile Gln
595 600 605

Glu Pro Thr Ala Asn Ala Thr Thr Thr Thr Ala Thr Thr Asp Glu Leu
610 615 620

Lys Thr Val Thr Lys Asp Arg Met Glu Asp Ile Lys Ile Leu Ile Ala
625 630 635 640

Ser Pro Ser Pro Thr His Ile His Lys Glu Thr Thr Ser Ala Thr Ser
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Ser Pro Tyr Arg Asp Thr Gln Ser Arg Thr Ala Ser Pro Asn Arg Ala
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Gly Lys Gly Val Ile Glu Gln Thr Glu Lys Ser His Pro Arg Ser Pro
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Asn Val Leu Ser Val Ala Leu Ser Gln Arg Thr Thr Val Pro Glu Glu
690 695 700

Glu Leu Asn Pro Lys Ile Leu Ala Leu Gln Asn Ala Gln Arg Lys Arg
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Lys Met Glu His Asp Gly Ser Leu Phe Gln Ala Val Gly Ile Gly Thr
725 730 735

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740 745 750

Lys Arg Val Lys Gly Cys Lys Ser Ser Glu Gln Asn Gly Met Glu Gln
755 760 765

Lys Thr Ile Ile Leu Ile Pro Ser Asp Leu Ala Cys Arg Leu Leu Gly
770 775 780

Gln Ser Met Asp Glu Ser Gly Leu Pro Gln Leu Thr Ser Tyr Asp Cys
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Ser Asp Met Val Pro Thr Cys Ser Ala Leu Ala Arg Lys Pro Asp Lys
115 120 125

Thr Gly Asn Thr Ser Thr Asp Gly Ser Tyr Lys Pro Ser Phe Leu Thr
145 150 155 160

Phe Ile Val Ser Cys Glu Thr Gly Arg Val Val Tyr Val Ser Asp Ser
180 185 190

Leu Tyr Asp Gln Val His Pro Asp Asp Val Asp Lys Leu Arg Glu Gln
210 215 220

Pro Tyr Ser Asp Glu Ile Glu Tyr Ile Ile Cys Thr Asn Thr Asn Val
450 455 460

Lys Asn Ser Ser Gln Glu Pro Arg Pro Thr Leu Ser Asn Thr Ile Gln
465 470 475 480

Arg Pro Gln Leu Gly Pro Thr Ala Asn Leu Pro Leu Glu Met Gly Ser
485 490 495

Gly Gln Leu Ala Pro Arg Gln Gln Gln Gln Thr Glu Leu Asp Met
500 505 510

Val Pro Gly Arg Asp Gly Leu Ala Ser Tyr Asn His Ser Gln Val Val
515 520 525

Gln Pro Val Thr Thr Thr Gly Pro Glu His Ser Lys Pro Leu Glu Lys
530 535 540

Ser Asp Gly Leu Phe Ala Gln Asp Arg Asp Pro Arg Phe Ser Glu Ile
545 550 555 560

Tyr His Asn Ile Asn Ala Asp Gln Ser Lys Gly Ile Ser Ser Ser Thr
565 570 575

Val Pro Ala Thr Gln Gln Leu Phe Ser Gln Gly Asn Thr Phe Pro Pro
580 585 590

Thr Pro Arg Pro Ala Glu Asn Phe Arg Asn Ser Gly Leu Ala Pro Pro
595 600 605

Val Thr Ile Val Gln Pro Ser Ala Ser Ala Gly Gln Met Leu Ala Gln
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Ile Ser Arg His Ser Asn Pro Thr Gln Gly Ala Thr Pro Thr Trp Thr
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Pro Thr Thr Arg Ser Gly Phe Ser Ala Gln Gln Val Ala Thr Gln Ala
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Thr Ala Lys Thr Arg Thr Ser Gln Phe Gly Val Gly Ser Phe Gln Thr
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Pro Ser Ser Phe Ser Ser Met Ser Leu Pro Gly Ala Pro Thr Ala Ser
675 680 685

Pro Gly Ala Ala Ala Tyr Pro Ser Leu Thr Asn Arg Gly Ser Asn Phe
690 695 700

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Ala Pro Glu Thr Gly Gln Thr Ala Gly Gln Phe Gln Thr Arg Thr Ala
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Glu Gly Val Gly Val Trp Pro Gln Trp Gln Gly Gln Gln Pro His His
725 730 735

Arg Ser Ser Ser Ser Glu Gln His Val Gln Gln Pro Pro Ala Gln Gln
740 745 750

Pro Gly Gln Pro Glu Val Phe Gln Glu Met Leu Ser Met Leu Gly Asp
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Gln Ser Asn Ser Tyr Asn Asn Glu Glu Phe Pro Asp Leu Thr Met Phe
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Pro Pro Phe Ser Glu
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Glu Ser Tyr Cys Gln Arg Gln Gly Val Pro Met Asn Ser Leu Arg Phe
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Leu Phe Glu Gly Gln Arg Ile Ala Asp Asn His Thr Pro Lys Glu Leu
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tggacaaggc ctccatcatg cgactggaaa tcagcttctt gcgaacacac aagctcctct	360
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<212> PRT
<213> homo sapiens

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<400> 60

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Glu Lys Ser Arg Asp Ala Ala Arg Cys Arg Arg Ser Lys Glu Thr Glu
20          25          30

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```

Val Phe Tyr Glu Leu Ala His Glu Leu Pro Leu Pro His Ser Val Ser

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35					40					45					
Ser	His	Leu	Asp	Lys	Ala	Ser	Ile	Met	Arg	Leu	Glu	Ile	Ser	Phe	Leu
50						55					60				
Arg	Thr	His	Lys	Leu	Leu	Ser	Ser	Val	Cys	Ser	Glu	Asn	Glu	Ser	Glu
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Ala	Glu	Ala	Asp	Gln	Gln	Met	Asp	Asn	Leu	Tyr	Leu	Lys	Ala	Leu	Glu
				85					90					95	
Gly	Phe	Ile	Ala	Val	Val	Thr	Gln	Asp	Gly	Asp	Met	Ile	Phe	Leu	Ser
			100					105					110		
Glu	Asn	Ile	Ser	Lys	Phe	Met	Gly	Leu	Thr	Gln	Val	Glu	Leu	Thr	Gly
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His	Ser	Ile	Phe	Asp	Phe	Thr	His	Pro	Cys	Asp	His	Glu	Glu	Ile	Arg
	130					135					140				
Glu	Asn	Leu	Ser	Leu	Lys	Asn	Gly	Ser	Gly	Phe	Gly	Lys	Lys	Ser	Lys
145					150					155					160
Asp	Met	Ser	Thr	Glu	Arg	Asp	Phe	Phe	Met	Arg	Met	Lys	Cys	Thr	Val
				165					170					175	
Thr	Asn	Arg	Gly	Arg	Thr	Val	Asn	Leu	Lys	Ser	Ala	Thr	Trp	Lys	Val
			180					185					190		
Leu	His	Cys	Thr	Gly	Gln	Val	Lys	Val	Tyr	Asn	Asn	Cys	Pro	Pro	His
		195					200					205			
Asn	Ser	Leu	Cys	Gly	Tyr	Lys	Glu	Pro	Leu	Leu	Ser	Cys	Leu	Ile	Ile
	210					215					220				
Met	Cys	Glu	Pro	Ile	Gln	His	Pro	Ser	His	Met	Asp	Ile	Pro	Leu	Asp
225					230					235					240
Ser	Lys	Thr	Phe	Leu	Ser	Arg	His	Ser	Met	Asp	Met	Lys	Phe	Thr	Tyr
				245					250					255	
Cys	Asp	Asp	Arg	Ile	Thr	Glu	Leu	Ile	Gly	Tyr	His	Pro	Glu	Glu	Leu
			260					265					270		
Leu	Gly	Arg	Ser	Ala	Tyr	Glu	Phe	Tyr	His	Ala	Leu	Asp	Ser	Glu	Asn

275	280	285
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Gly Gln Tyr Arg Met Leu Ala Lys His Gly Gly Tyr Val Trp Leu Glu 305 310 315 320		
Thr Gln Gly Thr Val Ile Tyr Asn Pro Arg Asn Leu Gln Pro Gln Cys 325 330 335		
Ile Met Cys Val Asn Tyr Val Leu Ser Glu Ile Glu Lys Asn Asp Val 340 345 350		
Val Phe Ser Met Asp Gln Thr Glu Ser Leu Phe Lys Pro His Leu Met 355 360 365		
Ala Met Asn Ser Ile Phe Asp Ser Ser Gly Lys Gly Ala Val Ser Glu 370 375 380		
Lys Ser Asn Phe Leu Phe Thr Lys Leu Lys Glu Glu Pro Glu Glu Leu 385 390 395 400		
Ala Gln Leu Ala Pro Thr Pro Gly Asp Ala Ile Ile Ser Leu Asp Phe 405 410 415		
Gly Asn Gln Asn Phe Glu Glu Ser Ser Ala Tyr Gly Lys Ala Ile Leu 420 425 430		
Pro Pro Ser Gln Pro Trp Ala Thr Glu Leu Arg Ser His Ser Thr Gln 435 440 445		
Ser Glu Ala Gly Ser Leu Pro Ala Phe Thr Val Pro Gln Ala Ala Ala 450 455 460		
Pro Gly Ser Thr Thr Pro Ser Ala Thr Ser Ser Ser Ser Cys Ser 465 470 475 480		
Thr Pro Asn Ser Pro Glu Asp Tyr Tyr Thr Ser Leu Asp Asn Asp Leu 485 490 495		
Lys Ile Glu Val Ile Glu Lys Leu Phe Ala Met Asp Thr Glu Ala Lys 500 505 510		
Asp Gln Cys Ser Thr Gln Thr Asp Phe Asn Glu Leu Asp Leu Glu Thr		

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515	520	525
Leu Ala Pro Tyr Ile Pro Met Asp Gly Glu Asp Phe Gln Leu Ser Pro 530 535 540		
Ile Cys Pro Glu Glu Arg Leu Leu Ala Glu Asn Pro Gln Ser Thr Pro 545 550 555 560		
Gln His Cys Phe Ser Ala Met Thr Asn Ile Phe Gln Pro Leu Ala Pro 565 570 575		
Val Ala Pro His Ser Pro Phe Leu Leu Asp Lys Phe Gln Gln Gln Leu 580 585 590		
Glu Ser Lys Lys Thr Glu Pro Glu His Arg Pro Met Ser Ser Ile Phe 595 600 605		
Phe Asp Ala Gly Ser Lys Ala Ser Leu Pro Pro Cys Cys Gly Gln Ala 610 615 620		
Ser Thr Pro Leu Ser Ser Met Gly Gly Arg Ser Asn Thr Gln Trp Pro 625 630 635 640		
Pro Asp Pro Pro Leu His Phe Gly Pro Thr Lys Trp Ala Val Gly Asp 645 650 655		
Gln Arg Thr Glu Phe Leu Gly Ala Ala Pro Leu Gly Pro Pro Val Ser 660 665 670		
Pro Pro His Val Ser Thr Phe Lys Thr Arg Ser Ala Lys Gly Phe Gly 675 680 685		
Ala Arg Gly Pro Asp Val Leu Ser Pro Ala Met Val Ala Leu Ser Asn 690 695 700		
Lys Leu Lys Leu Lys Arg Gln Leu Glu Tyr Glu Glu Gln Ala Phe Gln 705 710 715 720		
Asp Leu Ser Gly Gly Asp Pro Pro Gly Gly Ser Thr Ser His Leu Met 725 730 735		
Trp Lys Arg Met Lys Asn Leu Arg Gly Gly Ser Cys Pro Leu Met Pro 740 745 750		
Asp Lys Pro Leu Ser Ala Asn Val Pro Asn Asp Lys Phe Thr Gln Asn		

755	760	765
Pro Met Arg Gly Leu Gly His Pro Leu Arg His Leu Pro Leu Pro Gln		
770	775	780
Pro Pro Ser Ala Ile Ser Pro Gly Glu Asn Ser Lys Ser Arg Phe Pro		
785	790	800
Pro Gln Cys Tyr Ala Thr Gln Tyr Gln Asp Tyr Ser Leu Ser Ser Ala		
805	810	815
His Lys Val Ser Gly Met Ala Ser Arg Leu Leu Gly Pro Ser Phe Glu		
820	825	830
Ser Tyr Leu Leu Pro Glu Leu Thr Arg Tyr Asp Cys Glu Val Asn Val		
835	840	845
Pro Val Leu Gly Ser Ser Thr Leu Leu Gln Gly Gly Asp Leu Leu Arg		
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Ala Leu Asp Gln Ala Thr		
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 <212> DNA
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<400> 61

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gagctcattg gacacagcat ctttgatttc atccaccct gtgaccaaga ggagcttcag	420
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tgcttctcct tgcgcatgaa gactacgctc accagccgcg ggcgcaccct caacctcaag	540
gcggccacct ggaaggtgct gaactgctct ggacatatga gggcctacaa gccacctgcg	600
cagacttctc cagctgggag cctgactca gagccccgc tgcaagtgcct ggtgctcatc	660

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<400> 62

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Tyr Gln Leu Ala His Thr Leu Pro Phe Ala Arg Gly Val Ser Ala His
35 40 45

Leu Asp Lys Ala Ser Ile Met Arg Leu Thr Ile Ser Tyr Leu Arg Met
50 55 60

His Arg Leu Cys Ala Ala Gly Glu Trp Asn Gln Val Gly Ala Gly Gly
65 70 75 80

Glu Pro Leu Asp Ala Cys Tyr Leu Lys Ala Leu Glu Gly Phe Val Met
85 90 95

Val Leu Thr Ala Glu Gly Asp Met Ala Tyr Leu Ser Glu Asn Val Ser
100 105 110

Lys His Leu Gly Leu Ser Gln Leu Glu Leu Ile Gly His Ser Ile Phe
115 120 125

Asp Phe Ile His Pro Cys Asp Gln Glu Glu Leu Gln Asp Ala Leu Thr
130 135 140

Pro Gln Gln Thr Leu Ser Arg Arg Lys Val Glu Ala Pro Thr Glu Arg
145 150 155 160

Cys Phe Ser Leu Arg Met Lys Ser Thr Leu Thr Ser Arg Gly Arg Thr
165 170 175

Leu Asn Leu Lys Ala Ala Thr Trp Lys Val Leu Asn Cys Ser Gly His
180 185 190

Met Arg Ala Tyr Lys Pro Pro Ala Gln Thr Ser Pro Ala Gly Ser Pro
195 200 205

Asp Ser Glu Pro Pro Leu Gln Cys Leu Val Leu Ile Cys Glu Ala Ile
210 215 220

Pro His Pro Gly Ser Leu Glu Pro Pro Leu Gly Arg Gly Ala Phe Leu
225 230 235 240

Ser Arg His Ser Leu Asp Met Lys Phe Thr Tyr Cys Asp Asp Arg Ile
245 250 255

Ala Glu Val Ala Gly Tyr Ser Pro Asp Asp Leu Ile Gly Cys Ser Ala
260 265 270

Tyr Glu Tyr Ile His Ala Leu Asp Ser Asp Ala Val Ser Lys Ser Ile
275 280 285

His Thr Leu Leu Ser Lys Gly Gln Ala Val Thr Gly Gln Tyr Arg Phe
290 295 300

Leu Ala Arg Ser Gly Gly Tyr Leu Trp Thr Gln Thr Gln Ala Thr Val
305 310 315 320

Val Ser Gly Gly Arg Gly Pro Gln Ser Glu Ser Ile Val Cys Val His
325 330 335

Phe Leu Ile Ser Gln Val Glu Glu Thr Gly Val Val Leu Ser Leu Glu
340 345 350

Gln Thr Glu Gln His Ser Arg Arg Pro Ile Gln Arg Gly Ala Pro Ser
355 360 365

Gln Lys Asp Thr Pro Asn Pro Gly Asp Ser Leu Asp Thr Pro Gly Pro
370 375 380

Arg Ile Leu Ala Phe Leu His Pro Pro Ser Leu Ser Glu Ala Ala Leu
385 390 395 400

Ala Ala Asp Pro Arg Arg Phe Cys Ser Pro Asp Leu Arg Arg Leu Leu
405 410 415

Gly Pro Ile Leu Asp Gly Ala Ser Val Ala Ala Thr Pro Ser Thr Pro
420 425 430

Leu Ala Thr Arg His Pro Gln Ser Pro Leu Ser Ala Asp Leu Pro Asp
435 440 445

Glu Leu Pro Val Gly Thr Glu Asn Val His Arg Leu Phe Thr Ser Gly
450 455 460

Lys Asp Thr Glu Ala Val Glu Thr Asp Leu Asp Ile Ala Gln Asp Ala
465 470 475 480

Asp Ala Leu Asp Leu Glu Met Leu Ala Pro Tyr Ile Ser Met Asp Asp
485 490 495

Asp Phe Gln Leu Asn Ala Ser Glu Gln Leu Pro Arg Ala Tyr His Arg
500 505 510

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Pro Leu Gly Ala Val Pro Arg Pro Arg Ala Arg Ser Phe His Gly Leu
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Ser Pro Pro Ala Leu Glu Pro Ser Leu Leu Pro Arg Trp Gly Ser Asp
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Pro Arg Leu Ser Cys Ser Ser Pro Ser Arg Gly Asp Pro Ser Ala Ser
545 550 555 560

Ser Pro Met Ala Gly Ala Arg Lys Arg Thr Leu Ala Gln Ser Ser Glu
565 570 575

Asp Glu Asp Glu Gly Val Glu Leu Leu Gly Val Arg Pro Pro Lys Arg
580 585 590

Ser Pro Ser Pro Glu His Glu Asn Phe Leu Leu Phe Pro Leu Ser Leu
595 600 605

Ser Phe Leu Leu Thr Gly Gly Pro Ala Pro Gly Ser Leu Gln Asp Pro
610 615 620

Ser Thr Pro Leu Leu Asn Leu Asn Glu Pro Leu Gly Leu Gly Pro Ser
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Leu Leu Ser Pro Tyr Ser Asp Glu Asp Thr Thr Gln Pro Gly Gly Pro
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Phe Gln Pro Arg Ala Gly Ser Ala Gln Ala Asp
660 665

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Tyr Gly Arg Lys Lys Arg Arg Gln Arg Arg Arg Gly
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Ala Gln Arg Lys Arg Lys Met Glu His Asp Gly
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